

PATENT

C-3182/03

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

IN RE APPLICATION OF

SIKORSKI ET AL

GROUP ART UNIT: 1624

SERIAL NUMBER: NOT YET ASSIGNED

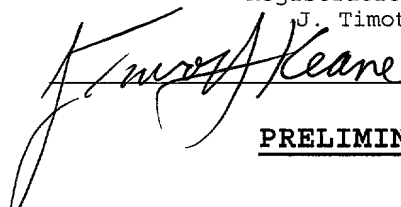
EXAMINER: RICHARD L. RAYMOND

FILED: 12 DEC 01

DATE: 12 DEC 01

TITLE: **(R)-CHIRAL HALOGENATED SUBSTITUTED N,N-Bis-PHENYL
AMINOALCOHOL COMPOUNDS USEFUL FOR INHIBITING
CHOLESTERYL ESTER TRANSFER PROTEIN ACTIVITY**

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Registration No. 27,808
J. Timothy Keane

 Date: 12 Dec 01

PRELIMINARY AMENDMENT

TO THE COMMISSIONER OF PATENTS AND TRADEMARKS

SIR:

Please amend the subject patent application as follows:

IN THE SPECIFICATION

Delete the originally-filed "Abstract" and replace with a
new -- Abstract --, attached hereto.

On page 1, delete the "Title", insert a new "Title" as
follows: -- (R)-CHIRAL HALOGENATED SUBSTITUTED N,N-Bis-PHENYL
AMINOALCOHOL COMPOUNDS USEFUL FOR INHIBITING CHOLESTERYL ESTER
TRANSFER PROTEIN ACTIVITY. --

On page 1, after the "Title", insert the following:

-- RELATED APPLICATIONS

This is a divisional of pending U.S. patent application
Serial No. 09/401,915, filed 23 September 1999. --

IN THE CLAIMS

Cancel Claims 1-67, without prejudice.

Add new Claims 68-98, attached hereto as Appendix A.

* * *

REMARKS

Originally-filed Claims 1-67 are now cancelled, without
prejudice, in favor of new Claims 68-98.

New Claims 68-98 are now in this application as the only
pending claims. These new claims are now introduced in response
to a Restriction Requirement originally imposed by the Patent
Office in parent Application Ser. No. 09/401,915 [Paper No. 8].

In Paper No. 8, Restriction has been imposed among the
following groups of subject matter:

Group I: Claims 1-67 (part), drawn to chiral benzyl phenyl
aminoalcohols, classified in class 514, subclass 655.

Group II: Claims 1-67 (part), drawn to chiral bis-benzyl aminoalcohols, classified in class 514, subclass 655.

Group III: Claims 1-67 (part), drawn to chiral bis-phenyl aminoalcohols, classified in class 514, subclass 658.

Group IV: Claims 1-67 (part), drawn to chiral phenoxy phenyl aminoalcohols, classified in class 514, subclass 645.

Group V: Claims 1-67 (part), drawn to chiral fused heterocyclic compounds, classified in class 514, subclass 230.5.

Group VI: Claims 1-67 (part), drawn to chiral heteroaryl benzyl aminoalcohols, classified in class 514, subclass 241.

In Applicants' response dated 29 October 2001, Applicants elected, with traverse, Group I subject matter. New independent generic Claims 68, 77 and 86 read on this elected species compound #8. This Group I subject matter is now covered in a divisional application to be co-filed with this present application. In this present application, new Claims 68-98 cover subject matter corresponding to previously non-elected Group III.

* * *

New Claims 68-98 define compound-per-se, composition and method-of-use subject matter containing chemical recitations consistent with Group III of Examiner's Restriction Requirement.

Support for new Claims 68-98 is found in originally-filed Claims 1-67. No new matter has been added.

* * *

The specification has been amended to include a new Title and Abstract consistent with the subject matter of new Claims 68-98.

Also, the specification has been amended at page 1 to identify the subject application as a divisional of parent Application Ser. No. 09/401,915 filed 23 September 1999.

* * *

In co-pending Application Ser. No. 09/405,524, the originally filed claims therein were rejected under 35 U.S.C. §112, second paragraph, as having improper dependencies among compound, composition and method-of-use claims [Paper No. 12 of U.S. Application Ser. No. 09/405,524]. The recent Office Action in the present application urges applicants to keep this §112 rejection in mind when filing any new claims in the present application.

Accordingly, new Claims 68-98 have been written to avoid these rejections under U.S.C. §112, second paragraph, and thus such §112 rejections should not be asserted against new Claims 68-98.

* * *

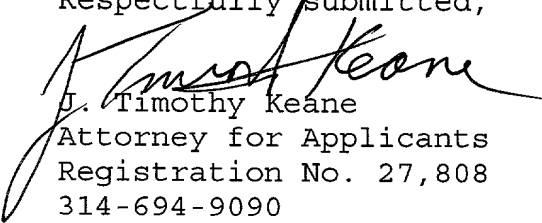
An Information Disclosure Statement under 37 CFR 1.97-1.99 will be filed along with references discussed with Examiner during a personal interview (4 April 2001) related to parent Application Ser. No. 09/401,915, shortly after Serial Number and

filing date information is received by applicants for the present divisional application.

* * *

In view of foregoing remarks, new Claims 68-98 should be in condition for allowance.

Respectfully submitted,


J. Timothy Keane

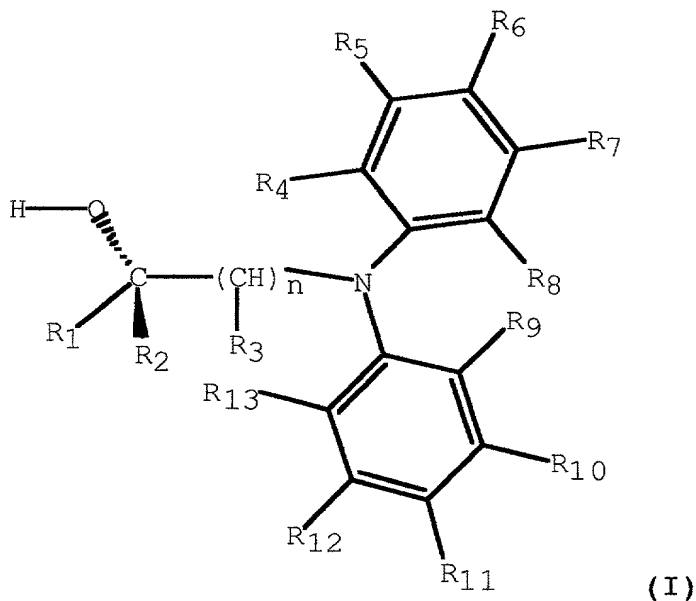
Attorney for Applicants
Registration No. 27,808
314-694-9090

Pharmacia Corporation
Corporate Patent Department
800 North Lindbergh Blvd.
Mail Zone O4E
St. Louis, MO 63167

Appendix A: New Claims 68-98

What we claim is:

68. A compound of Formula I:



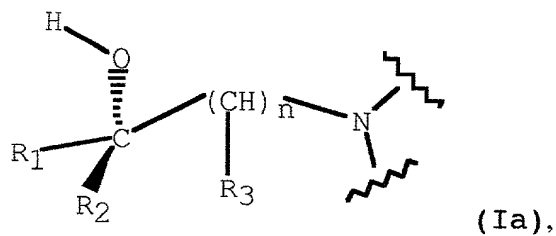
5

or a pharmaceutically acceptable salt thereof, wherein;

n is 1 or 2;

R_1 is haloalkyl or haloalkoxyalkyl with the proviso that R_1 is selected to have the highest Cahn-Ingold-Prelog stereochemical system ranking of three groups bonded to the hydroxy-substituted carbon to which R_1 and R_2 are attached in radical Ia:

10



which radical Ia is a fragment of Formula I;

R_2 is selected from the group consisting of hydrido, aryl, aralkyl, alkyl, alkenyl, alkenyloxyalkyl, haloalkyl, haloalkenyl, halocycloalkyl, haloalkoxyalkyl, haloalkenyloxyalkyl, halocycloalkoxy, halocycloalkoxyalkyl,

15

perhaloaryl, perhaloaralkyl, perhaloaryloxyalkyl, heteroaryl, dicyanoalkyl, and carboalkoxycyanoalkyl;

- R_3 is selected from the group consisting of hydrido, hydroxy, cyano, aryl, aralkyl, acyl, alkoxy, alkyl, alkenyl, alkoxyalkyl, heteroaryl, alkenyloxyalkyl, haloalkyl, haloalkenyl, haloalkoxy, haloalkoxyalkyl, haloalkenyloxyalkyl, monocyanoalkyl, dicyanoalkyl, carboxamido, and carboxamidoalkyl;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido, halo, haloalkyl, and alkyl;

- R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, perhaloaryloxy, alkanoylalkyl, alkanoylalkoxy, alkanoyloxy, N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclthio, hydroxyalkoxy, carboxamidoalkoxy, alkoxycarbonylalkoxy, alkoxycarbonylalkenyloxy, aralkanoylalkoxy, aralkenoyl, N-alkylcarboxamido, N-haloalkylcarboxamido, N-cycloalkylcarboxamido, N-arylcarboxamidoalkoxy, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl, carboxy, heteroaralkylthio, heteroaralkoxy, cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy, heterocyclloxy, aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocycl, haloalkylthio, alkanoyloxy, alkoxy, alkoxyalkyl, cycloalkoxy, cycloalkylalkoxy, hydroxy, amino, thio, nitro, alkylamino, alkylthio, arylamino, aralkylamino, arylthio, arylthioalkyl, alkylsulfonyl, alkylsulfonamido, monoarylamidosulfonyl, arylsulfonyl, heteroarylthio, heteroarylsulfonyl, heterocyclsulfonyl, heterocyclthio, alkanoyl, alkenoyl, aroyl, heteroaroyl, aralkanoyl, heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl, alkynyl, alkenyloxy, alkylenedioxy, haloalkylenedioxy, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl, hydroxyhaloalkoxy, hydroxyalkyl, aryl, aryloxy, aralkoxy, saturated heterocycl, heteroaryl, heteroaryloxy, heteroaryloxyalkyl, heteroaralkyl, arylalkenyl, carboalkoxy, alkoxycarboxamido, alkylamidocarbonylamido, arylamidocarbonylamido, carboalkoxyalkyl, carboalkoxyalkenyl, carboxamido, carboxamidoalkyl, and cyano;

R_5 and R_6 are optionally taken together to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 members, a partially

saturated heterocyclyl ring having 5 through 8 members, a heteroaryl ring having 5 or 6 members, and an aryl ring, wherein said cycloalkenyl ring, said partially saturated heterocyclyl ring, said heteroaryl ring, and said aryl are optionally substituted by one or more substituents selected from the group

5 consisting of R₁₀, R₁₁, and R₁₂;

R₁₀ and R₁₁ are optionally taken together to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 members, a partially saturated heterocyclyl ring having 5 through 8 members, a heteroaryl ring having 5 or 6 members, and an aryl ring, wherein said cycloalkenyl ring, said partially saturated heterocyclyl ring, said heteroaryl ring, and said aryl is optionally substituted by one or more substituents selected from the group consisting of R₅, R₆, and R₇;

with the proviso that the groups R₅ and R₆ and the groups R₁₀ and R₁₁ are not simultaneously taken together to form two rings;

15 with the further proviso that at least one of R₄, R₅, R₆, R₇, and R₈ is not hydrido or with the further proviso that at least one of R₉, R₁₀, R₁₁, R₁₂, and R₁₃ is not hydrido.

20 69. Compound of Claim 68 or a pharmaceutically acceptable salt thereof, wherein at least one of R₄, R₅, R₆, R₇, and R₈ is not hydrido and at least one of R₉, R₁₀, R₁₁, R₁₂, and R₁₃ is not hydrido.

25 70. Compound of Claim 69 or a pharmaceutically acceptable salt thereof, wherein;
n is 1 or 2;

R_1 is haloalkyl or haloalkoxyalkyl with the proviso that R_1 is selected to have the highest Cahn-Ingold-Prelog stereochemical system ranking of said three groups bonded to the hydroxy-substituted carbon to which R_1 and R_2 are attached in said fragment of the Formula I and with the further proviso that said

5 haloalkyl has two or more halo substituents;

R_2 is hydrido;

R_3 is hydrido;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the

10 group consisting of hydrido, perhaloaryloxy, N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclthio, hydroxyalkoxy, carboxamidoalkoxy, alkoxy-carbonylalkoxy, alkoxy-carbonylalkenyloxy, aralkanoylalkoxy, aralkenoyl, N-arylcarboxamidoalkoxy, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl, heteroaralkoxy, heterocyclloxy, aralkylaryl, aralkyl,

15 haloalkylthio, alkoxy, cycloalkoxy, cycloalkylalkoxy, alkylamino, alkylthio, arylamino, arylthio, arylsulfonyl, heteroarylthio, heteroarylsulfonyl, aroyl, alkyl, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl, aryl, aryloxy, aralkoxy, saturated heterocyclyl, heteroaryl, heteroaryloxyalkyl, and heteroaryloxy;

20 with the proviso that at least one of R_4 , R_5 , R_6 , R_7 , and R_8 is not hydrido and with the further proviso that at least one of R_9 , R_{10} , R_{11} , R_{12} , and R_{13} is not hydrido.

25 71. Compound of Claim 70 or a pharmaceutically acceptable salt thereof, wherein;

n is 1;

R_1 is selected from the group consisting of trifluoromethyl,

1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_2 is hydrido;

R_3 is hydrido;

5 R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

R_5 and R_{10} are independently selected from the group consisting of

- 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy,
- 10 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy,
- 15 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl, cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl, cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy,
- 20 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorobenzyloxy, 3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, difluoromethoxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 3,5-difluorobenzyloxy,
- 25 4-difluoromethoxybenzyloxy, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl, 1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy,
- 30 4-ethoxyphenoxy, 4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl, 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy, 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy,

- 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy,
 2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy,
 4-fluoro-2-trifluoromethylbenzyloxy, 4-fluoro-3-trifluoromethylbenzyloxy,
 2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy,
 5 2-fluorobenzyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
 3-iodobenzyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzyloxy,
 10 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl,
 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
 3-methoxycarbonylprop-2-en-yloxy, 4-methoxyphenyl,
 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy,
 15 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 4-methylthiophenoxy, 2-naphth-yloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl,
 pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 20 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino,
 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy, propoxy,
 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, sec-butyl,
 4-sec-butylphenoxy, tert -butoxy, 3-tert -butylphenoxy, 4-tert -butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 25 2-(5,6,7,8-tetrahydronaphth-yloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy,
 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl,
 30 3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy,
 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,
 35 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,

3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy,
 3-trifluoromethylthiobenzyloxy, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 5 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy,
 trifluoromethyl, and trifluoromethoxy;

R_7 and R_{12} are independently selected from the group consisting of
 hydrido, fluoro, and trifluoromethyl.

10

72. Compound of Claim 71 or a pharmaceutically acceptable salt
 thereof, wherein;

n is 1;

R_1 is selected from the group consisting of trifluoromethyl,
 15 difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_2 is hydrido;

R_3 is hydrido;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

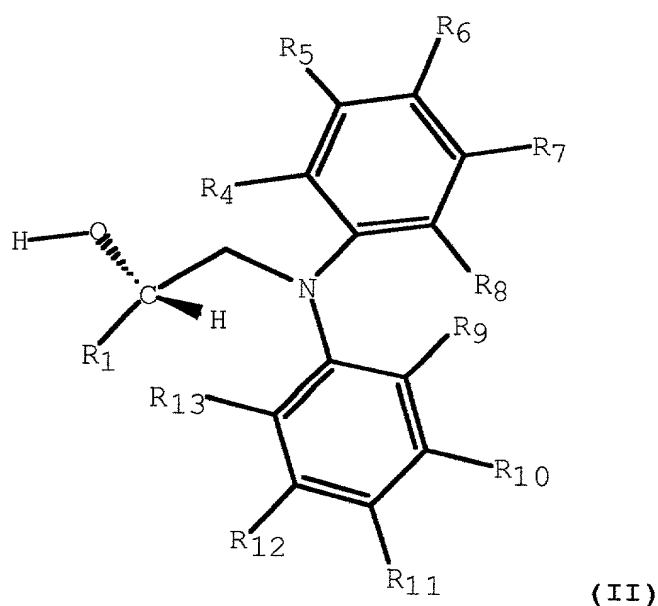
R_5 and R_{10} are independently selected from the group consisting of
 20 benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy,
 3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy, 3-chlorobenzyloxy,
 2-chlorophenoxy, 4-chloro-3-ethylphenoxy, 4-chloro-3-methylphenoxy,
 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy,
 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy,
 25 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy,
 4-chlorophenylamino, 5-chloropyrid-3-yloxy, cyclobutoxy, cyclobutyl,
 cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl,
 cyclopropylmethoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy,
 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl,
 30 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy,
 3,5-difluorobenzyloxy, difluoromethoxy, 3,5-difluorophenoxy,

- 3,4-difluorophenyl, 2,3-difluorophenoxy, 2,4-difluorophenoxy,
2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy,
3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 3,5-dimethylphenoxy,
3,4-dimethylphenoxy, 1,3-dioxolan-2-yl, 4-ethylbenzyloxy,
- 5 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy,
4-fluoro-3-methylbenzyl, 4-fluorobenzyloxy, 2-fluoro-3-methylphenoxy,
3-fluoro-4-methylphenoxy, 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy,
2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy,
2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy,
- 10 2-fluorobenzyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
2-furyl, 3-furyl, heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl,
2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl,
4-isoxazolyl, 5-isoxazolyl, isopropoxy, 4-isopropylbenzyloxy,
3-isopropylphenoxy, isopropylthio, 4-isopropyl-3-methylphenoxy,
- 15 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl, 3-methoxybenzyl,
4-methoxyphenylamino, 3-methylbenzyloxy, 4-methylbenzyloxy,
3-methylphenoxy, 3-methyl-4-methylthiophenoxy, 4-methylphenoxy,
1-methylpropoxy, 2-methylpyrid-5-yloxy, 4-methylthiophenoxy,
2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy, 3-nitrophenyl, 2-oxazolyl,
- 20 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio,
2,2,3,3,3-pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl,
1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy,
4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, tert-butoxy,
3-tert-butylphenoxy, 4-tert-butylphenoxy, 1,1,2,2-tetrafluoroethoxy,
- 25 tetrahydrofuran-2-yl, 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl,
thiazol-4-yl, thiazol-5-yl, thiophen-2-yl, 2,2,2-trifluoroethoxy,
2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
- 30 3-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy, 3-trifluoromethylphenyl,
2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
- 35 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy,
3-trifluoromethylthiobenzyloxy, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and trifluoromethyl;

R_7 and R_{12} are independently selected from the group consisting of
5 hydrido, fluoro, and trifluoromethyl.

73. Compound of Claim 70 of Formula II:



10 or a pharmaceutically acceptable salt thereof, wherein;

R_1 is haloalkyl;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the
group consisting of hydrido, perhaloaryloxy, N-aryl-N-alkylamino,
15 heterocyclalkoxy, heterocyclalthio, hydroxyalkoxy, aralkanoylalkoxy,
aralkenoyl, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl,
heteroaralkoxy, aralkyl, haloalkylthio, alkoxy, cycloalkoxy, cycloalkylalkoxy,
alkylthio, arylamino, arylthio, arylsulfonyl, aroyl, alkyl, cycloalkyl,
cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl,

hydroxyhaloalkoxy, aryl, aryloxy, aralkoxy, heteroaryl, heteroaryloxyalkyl, and heteroaryloxy;

with the proviso that at least one of R₄, R₅, R₆, R₇, and R₈ is not

hydrido and with the further proviso that at least one of R₉, R₁₀, R₁₁, R₁₂, and

5 R₁₃ is not hydrido.

74. Compound of Claim 73 or a pharmaceutically acceptable salt thereof, wherein;

10 R₁ is trifluoromethyl;

R₄, R₈, R₉, and R₁₃ are independently hydrido or fluoro;

R₅ is selected from the group consisting of 5-bromo-2-fluorophenoxy,

4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy,
3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy,

15 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy,
4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy,

3-pentafluoroethylphenoxy, 3-tert-butylphenoxy,
3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy),

20 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy,
3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R₁₀ is selected from the group consisting of cyclopentyl,

1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R₆, R₇, R₁₁, and R₁₂ are independently hydrido or fluoro.

25

75. Compound of Claim 74 or a pharmaceutically acceptable salt thereof, wherein;

R₁ is trifluoromethyl;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

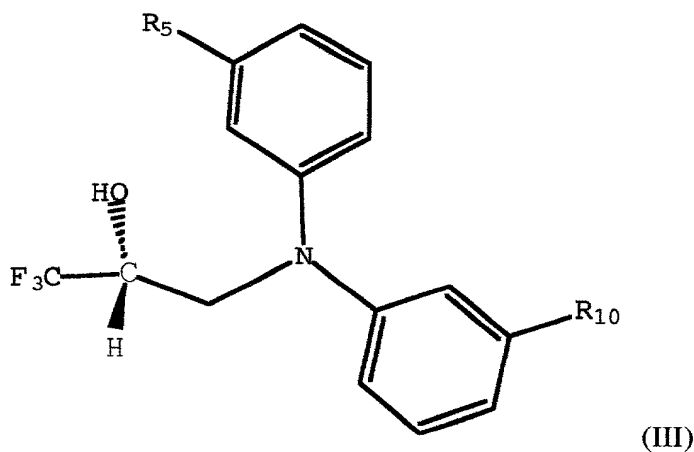
- R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 5 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy, 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 10 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of 1,1,2,2-tetrafluoroethoxy, pentafluoroethyl, and trifluoromethyl;

R_6 , R_7 , R_{11} , and R_{12} are independently hydrido or fluoro.

15

76. Compound of Claim 68 or a pharmaceutically acceptable salt thereof, wherein said compound is a compound of Formula III:



- wherein R_5 and R_{10} are selected from a compound selected from the group consisting of;
- 20

R_5 is 3-isopropylphenoxy and R_{10} is pentafluoroethyl;

R_5 is 2,3-dichlorophenoxy and R_{10} is pentafluoroethyl;

- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluorophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-methylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is pentafluoroethyl;
- 5 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-ethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-dimethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-*t*-butylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is pentafluoroethyl;
- 10 R_5 is 3,4-dichlorophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-dimethylaminophenoxy and R_{10} is pentafluoroethyl;
- 15 R_5 is 3-cyclopropylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-aminophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is pentafluoroethyl;
- 20 R_5 is 4-propoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is pentafluoroethyl;

- R_5 is 2-nitrophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is pentafluoroethyl;
- 5 R_5 is cyclohexylmethylenoxy and R_{10} is pentafluoroethyl;
- R_5 is benzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-ethylbenzyloxy and R_{10} is pentafluoroethyl;
- 10 R_5 is isopropoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is pentafluoroethyl;
- R_5 is isopropylthio and R_{10} is pentafluoroethyl;
- R_5 is cyclopentoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is pentafluoroethyl;
- 15 R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-isopropylbenzyloxy and R_{10} is pentafluoroethyl;
- 20 R_5 is 1-phenylethoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is pentafluoroethyl;

- R_5 is 3-trifluoromethylphenyl and R_{10} is pentafluoroethyl;
- R_5 is 4-methoxyphenylamino and R_{10} is pentafluoroethyl;
- R_5 is 4-nitrophenylthio and R_{10} is pentafluoroethyl;
- R_5 is 3-isopropylphenoxy and R_{10} is trifluoromethyl;
- 5 R_5 is 2,3-dichlorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-fluorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-methylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is trifluoromethyl;
- 10 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-ethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-dimethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-*t*-butylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is trifluoromethyl;
- 15 R_5 is 3,4-dichlorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is trifluoromethyl;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-dimethylaminophenoxy and R_{10} is trifluoromethyl;
- 20 R_5 is 3-cyclopropylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is trifluoromethyl;

- R_5 is 3-pentafluoroethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-aminophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-propoxyphenoxy and R_{10} is trifluoromethyl;
- 5 R_5 is 3-trifluoromethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 2-nitrophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is trifluoromethyl;
- 10 R_5 is cyclohexylmethylenoxy and R_{10} is trifluoromethyl;
- R_5 is benzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 4-ethylbenzyloxy and R_{10} is trifluoromethyl;
- 15 R_5 is isopropoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is trifluoromethyl;
- R_5 is isopropylthio and R_{10} is trifluoromethyl;
- R_5 is cyclopentoxy and R_{10} is trifluoromethyl;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is trifluoromethyl;
- 20 R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is trifluoromethyl;

- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 4-isopropylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 1-phenylethoxy and R_{10} is trifluoromethyl;
- 5 R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylphenyl and R_{10} is trifluoromethyl;
- R_5 is 4-methoxyphenylamino and R_{10} is trifluoromethyl;
- R_5 is 4-nitrophenylthio and R_{10} is trifluoromethyl;
- R_5 is 3-isopropylphenoxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 2,3-dichlorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-fluorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-methylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is trifluoromethoxy;
- 15 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-ethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-dimethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-*t*-butylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is trifluoromethoxy;
- 20 R_5 is 3,4-dichlorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is trifluoromethoxy;

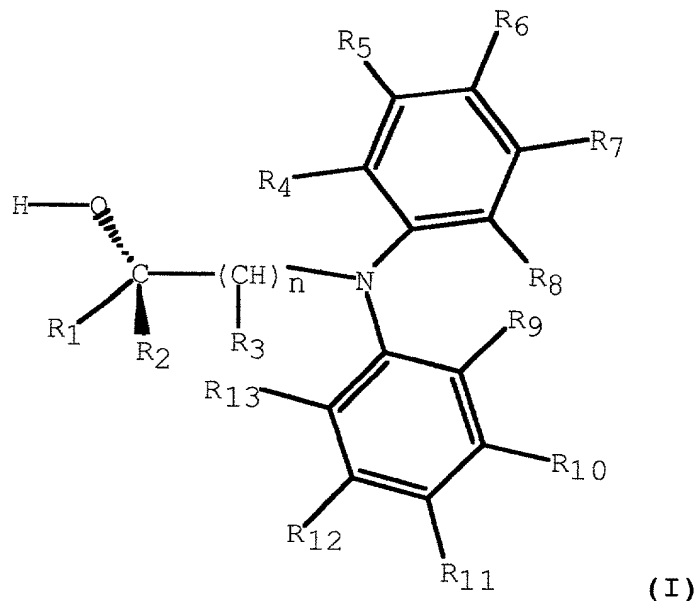
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-dimethylaminophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-cyclopropylphenoxy and R_{10} is trifluoromethoxy;
- 5 R_5 is 3-(2-furyl)phenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-aminophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-propoxyphenoxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 3-trifluoromethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 2-nitrophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is trifluoromethoxy;
- 15 R_5 is cyclohexylmethylenoxy and R_{10} is trifluoromethoxy;
- R_5 is benzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 4-ethylbenzyloxy and R_{10} is trifluoromethoxy;
- 20 R_5 is isopropoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is trifluoromethoxy;

- R_5 is isopropylthio and R_{10} is trifluoromethoxy;
- R_5 is cyclopentoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylthiobenzoyloxy and R_{10} is trifluoromethoxy;
- 5 R_5 is 3,4-dimethylbenzoyloxy and R_{10} is trifluoromethoxy;
- R_5 is 2-fluoro-3-trifluoromethylbenzoyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-fluoro-5-trifluoromethylbenzoyloxy and R_{10} is trifluoromethoxy;
- R_5 is 4-isopropylbenzoyloxy and R_{10} is trifluoromethoxy;
- R_5 is 1-phenylethoxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylphenyl and R_{10} is trifluoromethoxy;
- R_5 is 4-methoxyphenylamino and R_{10} is trifluoromethoxy;
- R_5 is 4-nitrophenylthio and R_{10} is trifluoromethoxy;
- R_5 is 3-isopropylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 2,3-dichlorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-methylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 20 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-ethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;

- R_5 is 3,5-dimethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-*t*-butylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,4-dichlorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 5 R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-dimethylaminophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 10 R_5 is 3-cyclopropylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-aminophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 4-propoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-nitrophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 20 R_5 is 3,5-difluorobenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is cyclohexylmethyleneoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is benzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;

- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-ethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is isopropoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 5 R_5 is 3-trifluoromethylbenzyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is isopropylthio and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is cyclopentoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 10 R_5 is 3,4-dimethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 4-isopropylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 1-phenylethoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylphenyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-methoxyphenylamino and R_{10} is 1,1,2,2-tetrafluoroethoxy; and
- 20 R_5 is 4-nitrophenylthio and R_{10} is 1,1,2,2-tetrafluoroethoxy.

77. A pharmaceutical composition comprising a therapeutically effective amount of a compound or a pharmaceutically acceptable salt thereof, together with a pharmaceutically acceptable carrier, said compound being of Formula I:

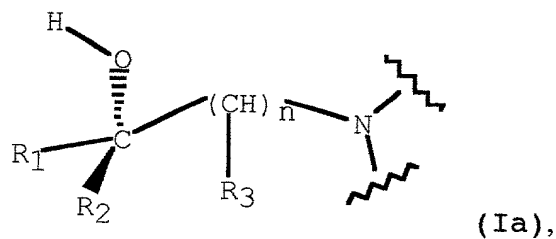


5 wherein;

n is 1 or 2;

R₁ is haloalkyl or haloalkoxyalkyl with the proviso that R₁ is selected to have the highest Cahn-Ingold-Prelog stereochemical system ranking of three groups bonded to the hydroxy-substituted carbon to which R₁ and R₂ are

10 attached in radical Ia:



which radical Ia is a fragment of Formula I;

R₂ is selected from the group consisting of hydrido, aryl, aralkyl, alkyl, alkenyl, alkenyloxyalkyl, haloalkyl, haloalkenyl, halocycloalkyl, haloalkoxyalkyl, haloalkenyloxyalkyl, halocycloalkoxy, halocycloalkoxyalkyl,

15

perhaloaryl, perhaloaralkyl, perhaloaryloxyalkyl, heteroaryl, dicyanoalkyl, and carboalkoxycyanoalkyl;

- R_3 is selected from the group consisting of hydrido, hydroxy, cyano, aryl, aralkyl, acyl, alkoxy, alkyl, alkenyl, alkoxyalkyl, heteroaryl,
- 5 alkenyloxyalkyl, haloalkyl, haloalkenyl, haloalkoxy, haloalkoxyalkyl, haloalkenyloxyalkyl, monocyanoalkyl, dicyanoalkyl, carboxamido, and carboxamidoalkyl;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido, halo, haloalkyl, and alkyl ;

- 10 R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, perhaloaryloxy, alkanoylalkyl, alkanoylalkoxy, alkanoyloxy, N-aryl-N-alkylamino, heterocyclylalkoxy, heterocyclylthio, hydroxyalkoxy, carboxamidoalkoxy, alkoxycarbonylalkoxy, alkoxycarbonylalkenyloxy, aralkanoylalkoxy, aralkenoyl,
- 15 N-alkylcarboxamido, N-haloalkylcarboxamido, N-cycloalkylcarboxamido, N-arylcarboxamidoalkoxy, cycloalkylcarbonyl, cyanoalkoxy, heterocyclylcarbonyl, carboxy, heteroaralkylthio, heteroaralkoxy, cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy, heterocycliloxy, aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocyclyl, haloalkylthio,
- 20 alkanoyloxy, alkoxy, alkoxyalkyl, cycloalkoxy, cycloalkylalkoxy, hydroxy, amino, thio, nitro, alkylamino, alkylthio, arylamino, aralkylamino, arylthio, arylthioalkyl, alkylsulfonyl, alkylsulfonamido, monoarylamidosulfonyl, arylsulfonyl, heteroarylthio, heteroarylsulfonyl, heterocyclylsulfonyl, heterocyclylthio, alkanoyl, alkenoyl, aroyl, heteroaroyl, aralkanoyl,
- 25 heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl, alkynyl, alkenyloxy, alkylenedioxy, haloalkylenedioxy, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl, hydroxyhaloalkoxy, hydroxyalkyl, aryl, aryloxy, aralkoxy, saturated heterocyclyl, heteroaryl, heteroaryloxy, heteroaryloxyalkyl, heteroaralkyl, arylalkenyl, carboalkoxy, alkoxycarboxamido,
- 30 alkylamidocarbonylamido, arylamidocarbonylamido, carboalkoxyalkyl, carboalkoxyalkenyl, carboxamido, carboxamidoalkyl, and cyano;

R_5 and R_6 are optionally taken together to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 members, a partially

saturated heterocyclyl ring having 5 through 8 members, a heteroaryl ring having 5 or 6 members, and an aryl ring, wherein said cycloalkenyl ring, said partially saturated heterocyclyl ring, said heteroaryl ring, and said aryl are optionally substituted by one or more substituents selected from the group

5 consisting of R_{10} , R_{11} , and R_{12} ;

R_{10} and R_{11} are optionally taken together to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 members, a partially saturated heterocyclyl ring having 5 through 8 members, a heteroaryl ring having 5 or 6 members, and an aryl ring, wherein said cycloalkenyl ring, said partially saturated heterocyclyl ring, said heteroaryl ring, and said aryl is optionally substituted by one or more substituents selected from the group consisting of R_5 , R_6 , and R_7 ;

with the proviso that the groups R_5 and R_6 and the groups R_{10} and R_{11} are not simultaneously taken together to form two rings;

15 with the further proviso that at least one of R_4 , R_5 , R_6 , R_7 , and R_8 is not hydrido or with the further proviso that at least one of R_9 , R_{10} , R_{11} , R_{12} , and R_{13} is not hydrido.

20 78. The pharmaceutical composition of Claim 77, wherein said compound is of Formula I, wherein at least one of R_4 , R_5 , R_6 , R_7 , and R_8 is not hydrido and at least one of R_9 , R_{10} , R_{11} , R_{12} , and R_{13} is not hydrido.

25 79. The pharmaceutical composition of Claim 78, wherein said compound is of Formula I, wherein;
n is 1 or 2;

R_1 is haloalkyl or haloalkoxyalkyl with the proviso that R_1 is selected to have the highest Cahn-Ingold-Prelog stereochemical system ranking of said three groups bonded to the hydroxy-substituted carbon to which R_1 and R_2 are attached in said fragment of the Formula I and with the further proviso that said

5 haloalkyl has two or more halo substituents;

R_2 is hydrido;

R_3 is hydrido;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the

10 group consisting of hydrido, perhaloaryloxy, N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclthio, hydroxyalkoxy, carboxamidoalkoxy, alkoxy-carbonylalkoxy, alkoxy-carbonylalkenyloxy, aralkanoylalkoxy, aralkenoyl, N-arylcarboxamidoalkoxy, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl, heteroaralkoxy, heterocyclloxy, aralkylaryl, aralkyl,

15 haloalkylthio, alkoxy, cycloalkoxy, cycloalkylalkoxy, alkylamino, alkylthio, arylamino, arylthio, arylsulfonyl, heteroarylthio, heteroarylsulfonyl, aroyl, alkyl, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl, hydroxyhaloalkoxy, aryl, aryloxy, aralkoxy, saturated heterocycl, heteroaryl, heteroaryloxyalkyl, and heteroaryloxy;

20 with the proviso that at least one of R_4 , R_5 , R_6 , R_7 , and R_8 is not hydrido and with the further proviso that at least one of R_9 , R_{10} , R_{11} , R_{12} , and R_{13} is not hydrido.

25 80. The pharmaceutical composition of Claim 79, wherein said compound is of Formula I, wherein;

n is 1;

R_1 is selected from the group consisting of trifluoromethyl, 1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_2 is hydrido;

5 R_3 is hydrido;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

R_5 and R_{10} are independently selected from the group consisting of

10 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy, 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy,

15 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl, cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl, cyclopentylcarbonyl,

20 cyclopropyl, cyclopropylmethoxy, cyclopropoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorobenzyloxy, 3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, difluoromethoxy, 3,5-difluorophenoxy,

25 3,4-difluorophenyl, 3,5-difluorobenzyloxy, 4-difluoromethoxybenzyloxy, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl, 1,4-dioxan-2-yl,

30 1,3-dioxolan-2-yl, ethoxy, 4-ethoxyphenoxy, 4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy, fluoro,

- 4-fluoro-3-methylbenzyl, 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy, 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy,
- 2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy,
- 5 4-fluoro-2-trifluoromethylbenzyloxy, 4-fluoro-3-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy, 2-fluorobenzyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy, 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
- 10 3-iodobenzyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl, 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzyloxy, 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio, 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl, 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
- 15 3-methoxycarbonylprop-2-en-yloxy, 4-methoxyphenyl, 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy, 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy, 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy, 4-methylthiophenoxy, 2-naphth-yloxy, 2-nitrophenoxy, 4-nitrophenoxy,
- 20 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy, propoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, sec-butyl,
- 25 4-sec-butylphenoxy, tert -butoxy, 3-tert -butylphenoxy, 4-tert -butylphenoxy, 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl, 2-(5,6,7,8-tetrahydronaphth-yloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl, thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
- 30 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl, 3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy, 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl, 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
- 35 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy, 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,

- 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy,
 5 3-trifluoromethylthiobenzyloxy, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy,
 trifluoromethyl, and trifluoromethoxy;

- R_7 and R_{12} are independently selected from the group consisting of
 10 hydrido, fluoro, and trifluoromethyl.

81. The pharmaceutical composition of Claim 80, wherein said
 compound is of Formula I, wherein;
 15 n is 1;

R_1 is selected from the group consisting of trifluoromethyl,
 difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_2 is hydrido;

R_3 is hydrido;

- 20 R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

- R_5 and R_{10} are independently selected from the group consisting of
 benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy,
 3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy, 3-chlorobenzyloxy,
 2-chlorophenoxy, 4-chloro-3-ethylphenoxy, 4-chloro-3-methylphenoxy,
 25 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy,
 3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy,
 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy,
 4-chlorophenylamino, 5-chloropyrid-3-yloxy, cyclobutoxy, cyclobutyl,
 cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl,
 30 cyclopropylmethoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy,
 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl,

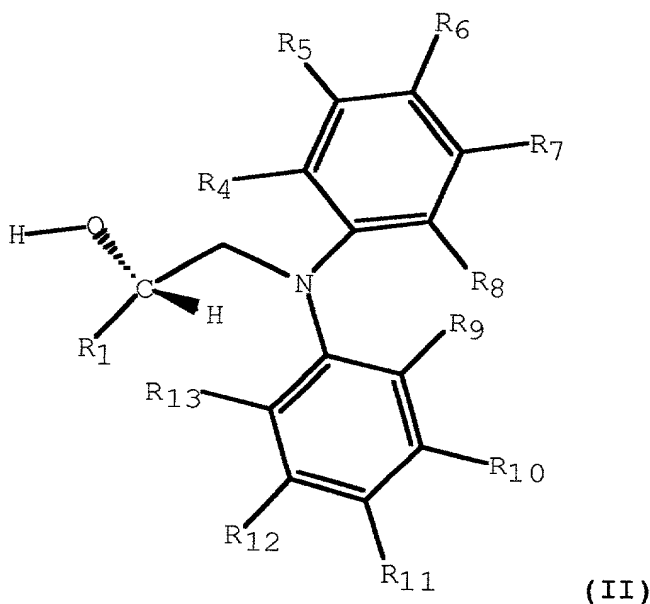
- 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzoyloxy,
 3,5-difluorobenzoyloxy, difluoromethoxy, 3,5-difluorophenoxy,
 3,4-difluorophenyl, 2,3-difluorophenoxy, 2,4-difluorophenoxy,
 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy,
 5 3,4-dimethylbenzoyloxy, 3,5-dimethylbenzoyloxy, 3,5-dimethylphenoxy,
 3,4-dimethylphenoxy, 1,3-dioxolan-2-yl, 4-ethylbenzoyloxy,
 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy,
 4-fluoro-3-methylbenzyl, 4-fluorobenzoyloxy, 2-fluoro-3-methylphenoxy,
 3-fluoro-4-methylphenoxy, 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy,
 10 2-fluoro-3-trifluoromethylbenzoyloxy, 3-fluoro-5-trifluoromethylbenzoyloxy,
 2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy,
 2-fluorobenzoyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 2-furyl, 3-furyl, heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl,
 2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl,
 15 4-isoxazolyl, 5-isoxazolyl, isopropoxy, 4-isopropylbenzoyloxy,
 3-isopropylphenoxy, isopropylthio, 4-isopropyl-3-methylphenoxy,
 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl, 3-methoxybenzyl,
 4-methoxyphenylamino, 3-methylbenzoyloxy, 4-methylbenzoyloxy,
 3-methylphenoxy, 3-methyl-4-methylthiophenoxy, 4-methylphenoxy,
 20 1-methylpropoxy, 2-methylpyrid-5-yloxy, 4-methylthiophenoxy,
 2-naphthyl, 2-nitrophenoxy, 4-nitrophenoxy, 3-nitrophenyl, 2-oxazolyl,
 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio,
 2,2,3,3,3-pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl,
 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy,
 25 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, tert-butoxy,
 3-tert-butylphenoxy, 4-tert-butylphenoxy, 1,1,2,2-tetrafluoroethoxy,
 tetrahydrofuran-2-yl, 2-(5,6,7,8-tetrahydronaphthyl), thiazol-2-yl,
 thiazol-4-yl, thiazol-5-yl, thiophen-2-yl, 2,2,2-trifluoroethoxy,
 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 30 3-trifluoromethoxybenzoyloxy, 4-trifluoromethoxybenzoyloxy,
 4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
 3-trifluoromethylbenzoyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzoyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy, 3-trifluoromethylphenyl,
 35 2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,
 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,

3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy,
3-trifluoromethylthiobenzyloxy, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and
5 trifluoromethyl;

R_7 and R_{12} are independently selected from the group consisting of
hydrido, fluoro, and trifluoromethyl.

10 82. The pharmaceutical composition of Claim 79, wherein said
compound is of Formula II:



wherein;

R_1 is haloalkyl;

15 R_4 , R_8 , R_9 , and R_{13} are independently hydrido or halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the
group consisting of hydrido, perhaloaryloxy, N-aryl-N-alkylamino,
heterocyclalkoxy, heterocyclthio, hydroxyalkoxy, aralkanoylalkoxy,
aralkenoyl, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl,
20 heteroaralkoxy, aralkyl, haloalkylthio, alkoxy, cycloalkoxy, cycloalkylalkoxy,

alkylthio, arylamino, arylthio, arylsulfonyl, aroyl, alkyl, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl, hydroxyhaloalkoxy, aryl, aryloxy, aralkoxy, heteroaryl, heteroaryloxyalkyl, and heteroaryloxy;

- 5 with the proviso that at least one of R₄, R₅, R₆, R₇, and R₈ is not hydrido and with the further proviso that at least one of R₉, R₁₀, R₁₁, R₁₂, and R₁₃ is not hydrido.

- 10 83. The pharmaceutical composition of Claim 82, wherein said compound is of Formula II, wherein;

R₁ is trifluoromethyl;

R₄, R₈, R₉, and R₁₃ are independently hydrido or fluoro;

R₅ is selected from the group consisting of 5-bromo-2-fluorophenoxy,

- 15 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy,
- 20 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy, 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R₁₀ is selected from the group consisting of cyclopentyl,

- 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
- 25 pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R₆, R₇, R₁₁, and R₁₂ are independently hydrido or fluoro.

84. The pharmaceutical composition of Claim 83, wherein said compound is of Formula II, wherein;

R_1 is trifluoromethyl;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

5 R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy,

4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy,
3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy,
3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy,
4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy,

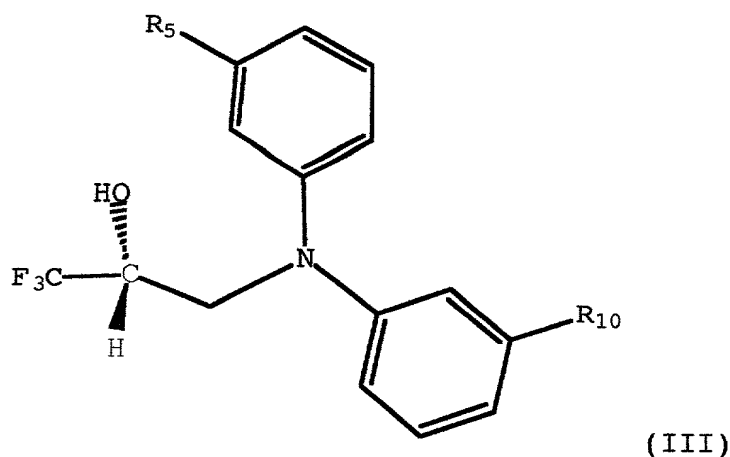
10 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy,
3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy,
3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy,
3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of 1,1,2,2-tetrafluoroethoxy,

15 pentafluoroethyl, and trifluoromethyl;

R_6 , R_7 , R_{11} , and R_{12} are independently hydrido or fluoro.

85. The pharmaceutical composition of Claim 77, wherein said
20 compound is a compound of Formula III:



wherein R_5 and R_{10} are selected to form a compound selected from the group consisting of;

- R_5 is 3-isopropylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 2,3-dichlorophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluorophenoxy and R_{10} is pentafluoroethyl;
- 5 R_5 is 4-methylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-chloro-3-ethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-ethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-dimethylphenoxy and R_{10} is pentafluoroethyl;
- 10 R_5 is 3-*t*-butylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3,4-dichlorophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is pentafluoroethyl;
- 15 R_5 is 3-difluoromethoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-dimethylaminophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-cyclopropylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is pentafluoroethyl;
- 20 R_5 is 4-aminophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is pentafluoroethyl;

- R_5 is 4-propoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 2-nitrophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is pentafluoroethyl;
- 5 R_5 is 3-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is cyclohexylmethylenoxy and R_{10} is pentafluoroethyl;
- R_5 is benzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- 10 R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-ethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is isopropoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is pentafluoroethyl;
- R_5 is isopropylthio and R_{10} is pentafluoroethyl;
- 15 R_5 is cyclopentoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- 20 R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-isopropylbenzyloxy and R_{10} is pentafluoroethyl;

- R_5 is 1-phenylethoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylphenyl and R_{10} is pentafluoroethyl;
- R_5 is 4-methoxyphenylamino and R_{10} is pentafluoroethyl;
- 5 R_5 is 4-nitrophenylthio and R_{10} is pentafluoroethyl;
- R_5 is 3-isopropylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 2,3-dichlorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-fluorophenoxy and R_{10} is trifluoromethyl;
- 10 R_5 is 4-methylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-chloro-3-ethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-ethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-dimethylphenoxy and R_{10} is trifluoromethyl;
- 15 R_5 is 3-*t*-butylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3,4-dichlorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is trifluoromethyl;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is trifluoromethyl;
- 20 R_5 is 3-difluoromethoxyphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-dimethylaminophenoxy and R_{10} is trifluoromethyl;

- R_5 is 3-cyclopropylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-aminophenoxy and R_{10} is trifluoromethyl;
- 5 R_5 is 3,4,5-trimethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-propoxyphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 2-nitrophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is trifluoromethyl;
- 10 R_5 is 3-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is trifluoromethyl;
- R_5 is cyclohexylmethylenoxy and R_{10} is trifluoromethyl;
- R_5 is benzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- 15 R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 4-ethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is isopropoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is trifluoromethyl;
- R_5 is isopropylthio and R_{10} is trifluoromethyl;
- 20 R_5 is cyclopentoxy and R_{10} is trifluoromethyl;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is trifluoromethyl;

- R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- 5 R_5 is 4-isopropylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 1-phenylethoxy and R_{10} is trifluoromethyl;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylphenyl and R_{10} is trifluoromethyl;
- R_5 is 4-methoxyphenylamino and R_{10} is trifluoromethyl;
- 10 R_5 is 4-nitrophenylthio and R_{10} is trifluoromethyl;
- R_5 is 3-isopropylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 2,3-dichlorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-fluorophenoxy and R_{10} is trifluoromethoxy;
- 15 R_5 is 4-methylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-chloro-3-ethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-ethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-dimethylphenoxy and R_{10} is trifluoromethoxy;
- 20 R_5 is 3-*t*-butylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is trifluoromethoxy;

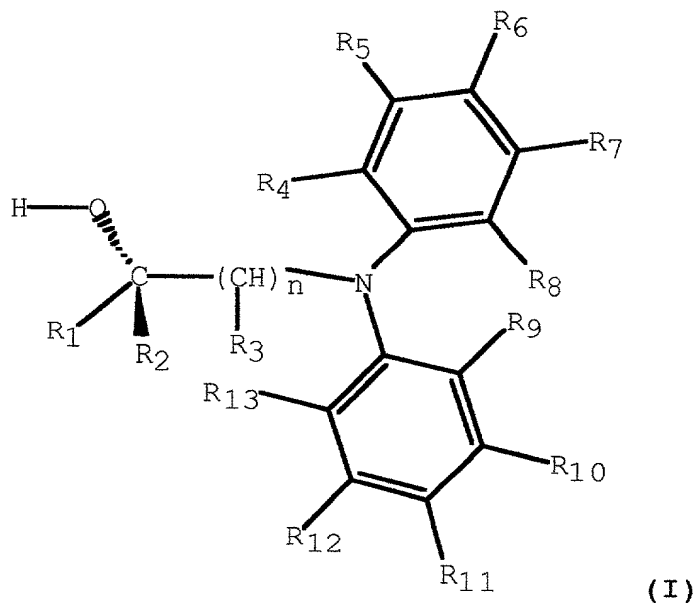
- R_5 is 3,4-dichlorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is trifluoromethoxy;
- 5 R_5 is 3-dimethylaminophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-cyclopropylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-aminophenoxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 3,4,5-trimethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-propoxyphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 2-nitrophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is trifluoromethoxy;
- 15 R_5 is 3-trifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is cyclohexylmethylenoxy and R_{10} is trifluoromethoxy;
- R_5 is benzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- 20 R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 4-ethylbenzyloxy and R_{10} is trifluoromethoxy;

- R_5 is isopropoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is trifluoromethoxy;
- R_5 is isopropylthio and R_{10} is trifluoromethoxy;
- R_5 is cyclopentoxy and R_{10} is trifluoromethoxy;
- 5 R_5 is 3-chloro-5-pyridinyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 4-isopropylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 1-phenylethoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylphenyl and R_{10} is trifluoromethoxy;
- R_5 is 4-methoxyphenylamino and R_{10} is trifluoromethoxy;
- 15 R_5 is 4-nitrophenylthio and R_{10} is trifluoromethoxy;
- R_5 is 3-isopropylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2,3-dichlorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 20 R_5 is 4-methylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;

- R_5 is 4-chloro-3-ethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-ethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,5-dimethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-*t*-butylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 5 R_5 is 4-fluoro-3-methylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,4-dichlorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 10 R_5 is 3-difluoromethoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-dimethylaminophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-cyclopropylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 4-aminophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-propoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-nitrophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 20 R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;

- R_5 is cyclohexylmethylenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is benzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 5 R_5 is 4-ethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is isopropoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is isopropylthio and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is cyclopentoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 10 R_5 is 3-chloro-5-pyridinyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-isopropylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 1-phenylethoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 20 R_5 is 3-trifluoromethylphenyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-methoxyphenylamino and R_{10} is 1,1,2,2-tetrafluoroethoxy; and
- R_5 is 4-nitrophenylthio and R_{10} is 1,1,2,2-tetrafluoroethoxy.

86. A method of treating or preventing a CETP-mediated disorder in a subject by administering a therapeutically effective amount of a compound or a pharmaceutically acceptable salt thereof, said compound being of Formula I:

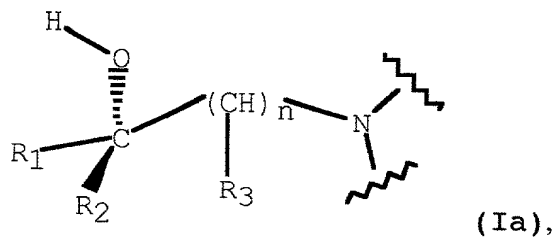


5 or a pharmaceutically acceptable salt thereof, wherein;

n is 1 or 2;

R₁ is haloalkyl or haloalkoxyalkyl with the proviso that R₁ is selected to have the highest Cahn-Ingold-Prelog stereochemical system ranking of three groups bonded to the hydroxy-substituted carbon to which R₁ and R₂ are

10 attached in radical Ia:



which radical Ia is a fragment of Formula I;

R₂ is selected from the group consisting of hydrido, aryl, aralkyl, alkyl, alkenyl, alkenyloxyalkyl, haloalkyl, haloalkenyl, halocycloalkyl, haloalkoxyalkyl, haloalkenyloxyalkyl, halocycloalkoxy, halocycloalkoxyalkyl,

15

perhaloaryl, perhaloaralkyl, perhaloaryloxyalkyl, heteroaryl, dicyanoalkyl, and carboalkoxycyanoalkyl;

- R_3 is selected from the group consisting of hydrido, hydroxy, cyano, aryl, aralkyl, acyl, alkoxy, alkyl, alkenyl, alkoxyalkyl, heteroaryl,
- 5 alkenyloxyalkyl, haloalkyl, haloalkenyl, haloalkoxy, haloalkoxyalkyl, haloalkenyloxyalkyl, monocyanoalkyl, dicyanoalkyl, carboxamido, and carboxamidoalkyl;

R_4 , R_8 , R_9 , and R_{13} are independently selected from the group consisting of hydrido, halo, haloalkyl, and alkyl ;

- 10 R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, perhaloaryloxy, alkanoylalkyl, alkanoylalkoxy, alkanoyloxy, N-aryl-N-alkylamino, heterocyclylalkoxy, heterocyclylthio, hydroxyalkoxy, carboxamidoalkoxy, alkoxycarbonylalkoxy, alkoxycarbonylalkenyloxy, aralkanoylalkoxy, aralkenoyl,
- 15 N-alkylcarboxamido, N-haloalkylcarboxamido, N-cycloalkylcarboxamido, N-arylcarboxamidoalkoxy, cycloalkylcarbonyl, cyanoalkoxy, heterocyclylcarbonyl, carboxy, heteroaralkylthio, heteroaralkoxy, cycloalkylamino, acylalkyl, acylalkoxy, aroylalkoxy, heterocycliloxy, aralkylaryl, aralkyl, aralkenyl, aralkynyl, heterocyclyl, haloalkylthio,
- 20 alkanoyloxy, alkoxy, alkoxyalkyl, cycloalkoxy, cycloalkylalkoxy, hydroxy, amino, thio, nitro, alkylamino, alkylthio, arylamino, aralkylamino, arylthio, arylthioalkyl, alkylsulfonyl, alkylsulfonamido, monoarylamidosulfonyl, arylsulfonyl, heteroarylthio, heteroarylsulfonyl, heterocyclylsulfonyl, heterocyclylthio, alkanoyl, alkenoyl, aroyl, heteroaroyl, aralkanoyl,
- 25 heteroaralkanoyl, haloalkanoyl, alkyl, alkenyl, alkynyl, alkenyloxy, alkylenedioxy, haloalkylenedioxy, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl, hydroxyhaloalkoxy, hydroxyalkyl, aryl, aryloxy, aralkoxy, saturated heterocyclyl, heteroaryl, heteroaryloxy, heteroaryloxyalkyl, heteroaralkyl, arylalkenyl, carboalkoxy, alkoxycarboxamido,
- 30 alkylamidocarbonylamido, arylamidocarbonylamido, carboalkoxyalkyl, carboalkoxyalkenyl, carboxamido, carboxamidoalkyl, and cyano;

R_5 and R_6 are optionally taken together to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 members, a partially

saturated heterocyclyl ring having 5 through 8 members, a heteroaryl ring having 5 or 6 members, and an aryl ring, wherein said cycloalkenyl ring, said partially saturated heterocyclyl ring, said heteroaryl ring, and said aryl are optionally substituted by one or more substituents selected from the group

- 5 consisting of R₁₀, R₁₁, and R₁₂;

- R₁₀ and R₁₁ are optionally taken together to form a ring selected from the group consisting of a cycloalkenyl ring having 5 through 8 members, a partially saturated heterocyclyl ring having 5 through 8 members, a heteroaryl ring having 5 or 6 members, and an aryl ring, wherein said cycloalkenyl ring, said partially saturated heterocyclyl ring, said heteroaryl ring, and said aryl is optionally substituted by one or more substituents selected from the group consisting of R₅, R₆, and R₇;
- 10

with the proviso that the groups R₅ and R₆ and the groups R₁₀ and R₁₁ are not simultaneously taken together to form two rings;

- 15 with the further proviso that at least one of R₄, R₅, R₆, R₇, and R₈ is not hydrido or with the further proviso that at least one of R₉, R₁₀, R₁₁, R₁₂, and R₁₃ is not hydrido.

- 20 87. The method of Claim 86, wherein said compound is of Formula I, wherein at least one of R₄, R₅, R₆, R₇, and R₈ that is not hydrido and at least one of R₉, R₁₀, R₁₁, R₁₂, and R₁₃ that is not hydrido.

- 25 88. The method of Claim 87, wherein said compound is of Formula I, wherein;
n is 1 or 2;

R_1 is haloalkyl or haloalkoxyalkyl with the proviso that R_1 is selected to have the highest Cahn-Ingold-Prelog stereochemical system ranking of said three groups bonded to the hydroxy-substituted carbon to which R_1 and R_2 are attached in said fragment of the Formula I and with the further proviso that said

5 haloalkyl has two or more halo substituents;

R_2 is hydrido;

R_3 is hydrido;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the

10 group consisting of hydrido, perhaloaryloxy, N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclthio, hydroxyalkoxy, carboxamidoalkoxy, alkoxy-carbonylalkoxy, alkoxy-carbonylalkenyloxy, aralkanoylalkoxy, aralkanoyl, N-arylcarboxamidoalkoxy, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl, heteroaralkoxy, heterocyclloxy, aralkylaryl, aralkyl,

15 haloalkylthio, alkoxy, cycloalkoxy, cycloalkylalkoxy, alkylamino, alkylthio, arylamino, arylthio, arylsulfonyl, heteroarylthio, heteroarylsulfonyl, aroyl, alkyl, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl, hydroxyhaloalkoxy, aryl, aryloxy, aralkoxy, saturated heterocycl, heteroaryl, heteroaryloxyalkyl, and heteroaryloxy;

20 with the proviso that at least one of R_4 , R_5 , R_6 , R_7 , and R_8 is not hydrido and with the further proviso that at least one of R_9 , R_{10} , R_{11} , R_{12} , and R_{13} is not hydrido.

25 89. The method of Claim 88, wherein said compound is of Formula I, wherein;

n is 1;

R_1 is selected from the group consisting of trifluoromethyl,

1,1,2,2-tetrafluoroethoxymethyl, trifluoromethoxymethyl, difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_2 is hydrido;

R_3 is hydrido;

5 R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

R_5 and R_{10} are independently selected from the group consisting of

- 4-aminophenoxy, benzoyl, benzyl, benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy, 4-bromo-2-nitrophenoxy, 3-bromobenzyloxy, 4-bromobenzyloxy, 4-bromophenoxy, 5-bromopyrid-2-yloxy,
- 10 4-butoxyphenoxy, chloro, 3-chlorobenzyl, 2-chlorophenoxy, 4-chlorophenoxy, 4-chloro-3-ethylphenoxy, 3-chloro-4-fluorobenzyl, 3-chloro-4-fluorophenyl, 3-chloro-2-fluorobenzyloxy, 3-chlorobenzyloxy, 4-chlorobenzyloxy, 4-chloro-3-methylphenoxy, 2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy, 3-chloro-4-ethylphenoxy,
- 15 3-chloro-4-methylphenoxy, 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy, 4-chlorophenylamino, 5-chloropyrid-3-yloxy, 2-cyanopyrid-3-yloxy, 4-cyanophenoxy, cyclobutoxy, cyclobutyl, cyclohexoxy, cyclohexylmethoxy, cyclopentoxy, cyclopentyl, cyclopentylcarbonyl, cyclopropyl, cyclopropylmethoxy, cyclopropoxy,
- 20 2,3-dichlorophenoxy, 2,4-dichlorophenoxy, 2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl, 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy, 2,4-difluorophenoxy, 3,4-difluorobenzyloxy, 2,5-difluorobenzyloxy, difluoromethoxy, 3,5-difluorophenoxy, 3,4-difluorophenyl, 3,5-difluorobenzyloxy,
- 25 4-difluoromethoxybenzyloxy, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 3,4-dimethylbenzyl, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 2,2-dimethylpropoxy, 1,3-dioxan-2-yl, 1,4-dioxan-2-yl, 1,3-dioxolan-2-yl, ethoxy,
- 30 4-ethoxyphenoxy, 4-ethylbenzyloxy, 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy, fluoro, 4-fluoro-3-methylbenzyl, 4-fluoro-3-methylphenyl, 4-fluoro-3-methylbenzoyl, 4-fluorobenzyloxy, 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy,

- 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy,
 2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy,
 4-fluoro-2-trifluoromethylbenzyloxy, 4-fluoro-3-trifluoromethylbenzyloxy,
 2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy,
 5 2-fluorobenzyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy,
 4-fluoropyrid-2-yloxy, 2-furyl, 3-furyl, heptafluoropropyl,
 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy,
 3-iodobenzyloxy, isobutyl, isobutylamino, isobutoxy, 3-isoxazolyl,
 4-isoxazolyl, 5-isoxazolyl, isopropoxy, isopropyl, 4-isopropylbenzyloxy,
 10 3-isopropylphenoxy, 4-isopropylphenoxy, isopropylthio,
 4-isopropyl-3-methylphenoxy, 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl,
 3-methoxybenzyl, 4-methoxycarbonylbutoxy,
 3-methoxycarbonylprop-2-enyloxy, 4-methoxyphenyl,
 3-methoxyphenylamino, 4-methoxyphenylamino, 3-methylbenzyloxy,
 15 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy,
 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy,
 4-methylthiophenoxy, 2-naphthylloxy, 2-nitrophenoxy, 4-nitrophenoxy,
 3-nitrophenyl, 4-nitrophenylthio, 2-oxazolyl, 4-oxazolyl, 5-oxazolyl,
 pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl,
 20 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino,
 1-phenylethoxy, phenylsulfonyl, 4-propanoylphenoxy, propoxy,
 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, sec-butyl,
 4-sec-butylphenoxy, tert-butoxy, 3-tert-butylphenoxy, 4-tert-butylphenoxy,
 1,1,2,2-tetrafluoroethoxy, tetrahydrofuran-2-yl,
 25 2-(5,6,7,8-tetrahydronaphthylloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl,
 thiophen-2-yl, 2,3,5-trifluorobenzyloxy, 2,2,2-trifluoroethoxy,
 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy,
 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy,
 3-trifluoromethoxyphenoxy, 4-trifluoromethoxyphenoxy, trifluoromethyl,
 30 3-trifluoromethylbenzyloxy, 4-trifluoromethylbenzyloxy,
 2,4-bis-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl,
 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy,
 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy,
 3-trifluoromethylphenyl, 3-trifluoromethylthiobenzyloxy,
 35 4-trifluoromethylthiobenzyloxy, 2,3,4-trifluorophenoxy,
 2,3,4-trifluorophenyl, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy,

3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy,
3-trifluoromethylthiobenzyloxy, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of
5 chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy,
trifluoromethyl, and trifluoromethoxy;

R_7 and R_{12} are independently selected from the group consisting of
hydrido, fluoro, and trifluoromethyl.

10

90. The method of Claim 89, wherein said compound is of Formula
I, wherein;

n is 1;

R_1 is selected from the group consisting of trifluoromethyl,
15 difluoromethyl, chlorodifluoromethyl, and pentafluoroethyl;

R_2 is hydrido;

R_3 is hydrido;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

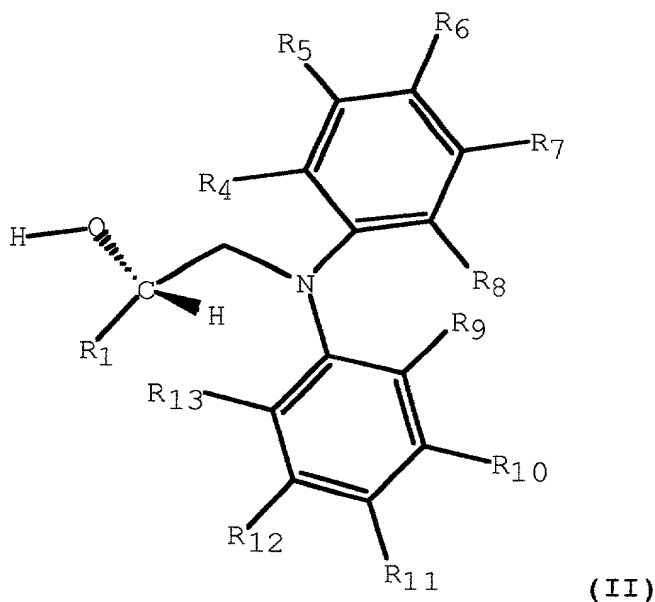
R_5 and R_{10} are independently selected from the group consisting of
20 benzyloxy, 5-bromo-2-fluorophenoxy, 4-bromo-3-fluorophenoxy,
3-bromobenzyloxy, 4-bromophenoxy, 4-butoxyphenoxy, 3-chlorobenzyloxy,
2-chlorophenoxy, 4-chloro-3-ethylphenoxy, 4-chloro-3-methylphenoxy,
2-chloro-4-fluorophenoxy, 4-chloro-2-fluorophenoxy, 4-chlorophenoxy,
3-chloro-4-ethylphenoxy, 3-chloro-4-methylphenoxy,
25 3-chloro-4-fluorophenoxy, 4-chloro-3-fluorophenoxy,
4-chlorophenylamino, 5-chloropyrid-3-yloxy, cyclobutoxy, cyclobutyl,
cyclohexylmethoxy, cyclopentoxo, cyclopentyl, cyclopentylcarbonyl,
cyclopropylmethoxy, 2,3-dichlorophenoxy, 2,4-dichlorophenoxy,
2,4-dichlorophenyl, 3,5-dichlorophenyl, 3,5-dichlorobenzyl,
30 3,4-dichlorophenoxy, 3,4-difluorophenoxy, 2,3-difluorobenzyloxy,
3,5-difluorobenzyloxy, difluoromethoxy, 3,5-difluorophenoxy,

- 3,4-difluorophenyl, 2,3-difluorophenoxy, 2,4-difluorophenoxy, 2,5-difluorophenoxy, 3,5-dimethoxyphenoxy, 3-dimethylaminophenoxy, 3,4-dimethylbenzyloxy, 3,5-dimethylbenzyloxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 1,3-dioxolan-2-yl, 4-ethylbenzyloxy,
- 5 3-ethylphenoxy, 4-ethylaminophenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylbenzyl, 4-fluorobenzyloxy, 2-fluoro-3-methylphenoxy, 3-fluoro-4-methylphenoxy, 3-fluorophenoxy, 3-fluoro-2-nitrophenoxy, 2-fluoro-3-trifluoromethylbenzyloxy, 3-fluoro-5-trifluoromethylbenzyloxy, 2-fluorophenoxy, 4-fluorophenoxy, 2-fluoro-3-trifluoromethylphenoxy,
- 10 2-fluorobenzyloxy, 4-fluorophenylamino, 2-fluoro-4-trifluoromethylphenoxy, 2-furyl, 3-furyl, heptafluoropropyl, 1,1,1,3,3,3-hexafluoropropyl, 2-hydroxy-3,3,3-trifluoropropoxy, isobutoxy, isobutyl, 3-isoxazolyl, 4-isoxazolyl, 5-isoxazolyl, isopropoxy, 4-isopropylbenzyloxy, 3-isopropylphenoxy, isopropylthio, 4-isopropyl-3-methylphenoxy,
- 15 3-isothiazolyl, 4-isothiazolyl, 5-isothiazolyl, 3-methoxybenzyl, 4-methoxyphenylamino, 3-methylbenzyloxy, 4-methylbenzyloxy, 3-methylphenoxy, 3-methyl-4-methylthiophenoxy, 4-methylphenoxy, 1-methylpropoxy, 2-methylpyrid-5-yloxy, 4-methylthiophenoxy, 2-naphthyloxy, 2-nitrophenoxy, 4-nitrophenoxy, 3-nitrophenyl, 2-oxazolyl,
- 20 4-oxazolyl, 5-oxazolyl, pentafluoroethyl, pentafluoroethylthio, 2,2,3,3,3-pentafluoropropyl, 1,1,3,3,3-pentafluoropropyl, 1,1,2,2,3-pentafluoropropyl, phenoxy, phenylamino, 1-phenylethoxy, 4-propylphenoxy, 4-propoxyphenoxy, thiophen-3-yl, tert-butoxy, 3-tert-butylphenoxy, 4-tert-butylphenoxy, 1,1,2,2-tetrafluoroethoxy,
- 25 tetrahydrofuran-2-yl, 2-(5,6,7,8-tetrahydronaphthyloxy), thiazol-2-yl, thiazol-4-yl, thiazol-5-yl, thiophen-2-yl, 2,2,2-trifluoroethoxy, 2,2,2-trifluoroethyl, 3,3,3-trifluoro-2-hydroxypropyl, trifluoromethoxy, 3-trifluoromethoxybenzyloxy, 4-trifluoromethoxybenzyloxy, 4-trifluoromethoxyphenoxy, 3-trifluoromethoxyphenoxy, trifluoromethyl,
- 30 3-trifluoromethylbenzyloxy, 1,1-bis-trifluoromethyl-1-hydroxymethyl, 3-trifluoromethylbenzyl, 3,5-bis-trifluoromethylbenzyloxy, 4-trifluoromethylphenoxy, 3-trifluoromethylphenoxy, 3-trifluoromethylphenyl, 2,3,4-trifluorophenoxy, 2,3,5-trifluorophenoxy, 3,4,5-trimethylphenoxy, 3-difluoromethoxyphenoxy, 3-pentafluoroethylphenoxy,
- 35 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 3-trifluoromethylthiophenoxy, 3-trifluoromethylthiobenzyloxy, and trifluoromethylthio;

R_6 and R_{11} are independently selected from the group consisting of chloro, fluoro, hydrido, pentafluoroethyl, 1,1,2,2-tetrafluoroethoxy, and trifluoromethyl;

R_7 and R_{12} are independently selected from the group consisting of hydrido, fluoro, and trifluoromethyl.

91. The method of Claim 88, wherein said compound is of Formula II:



10

wherein;

R_1 is haloalkyl;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or halo;

R_5 , R_6 , R_7 , R_{10} , R_{11} , and R_{12} are independently selected from the group consisting of hydrido, perhaloaryloxy, N-aryl-N-alkylamino, heterocyclalkoxy, heterocyclalthio, hydroxyalkoxy, aralkanoylalkoxy, aralkenoyl, cycloalkylcarbonyl, cyanoalkoxy, heterocyclcarbonyl, heteroaralkoxy, aralkyl, haloalkylthio, alkoxy, cycloalkoxy, cycloalkylalkoxy, alkylthio, arylamino, arylthio, arylsulfonyl, aroyl, alkyl, cycloalkyl, cycloalkylalkanoyl, halo, haloalkyl, haloalkoxy, hydroxyhaloalkyl,

20

hydroxyhaloalkoxy, aryl, aryloxy, aralkoxy, heteroaryl, heteroaryloxyalkyl, and heteroaryloxy;

with the proviso that at least one of R_4 , R_5 , R_6 , R_7 , and R_8 is not

hydrido and with the further proviso that at least one of R_9 , R_{10} , R_{11} , R_{12} , and

5 R_{13} is not hydrido.

92. The method of Claim 91, wherein said compound is of Formula II, wherein;

10 R_1 is trifluoromethyl;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy, 4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy, 3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy, 15 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy, 4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy, 3-pentafluoroethylphenoxy, 3-tert-butylphenoxy, 3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy, 3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy, 20 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

R_{10} is selected from the group consisting of cyclopentyl, 1,1,2,2-tetrafluoroethoxy, 2-furyl, 1,1-bis-trifluoromethyl-1-hydroxymethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethyl, and trifluoromethylthio;

R_6 , R_7 , R_{11} , and R_{12} are independently hydrido or fluoro.

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93. The method of Claim 92, wherein said compound is of Formula II, wherein;

R_1 is trifluoromethyl;

R_4 , R_8 , R_9 , and R_{13} are independently hydrido or fluoro;

R_5 is selected from the group consisting of 5-bromo-2-fluorophenoxy,

4-chloro-3-ethylphenoxy, 2,3-dichlorophenoxy, 3,4-dichlorophenoxy,

3-difluoromethoxyphenoxy, 3,5-dimethylphenoxy, 3,4-dimethylphenoxy,

5 3-ethylphenoxy, 3-ethyl-5-methylphenoxy, 4-fluoro-3-methylphenoxy,

4-fluorophenoxy, 3-isopropylphenoxy, 3-methylphenoxy,

3-pentafluoroethylphenoxy, 3-tert-butylphenoxy,

3-(1,1,2,2-tetrafluoroethoxy)phenoxy, 2-(5,6,7,8-tetrahydronaphthyl)oxy,

3-trifluoromethoxybenzyloxy, 3-trifluoromethoxyphenoxy,

10 3-trifluoromethylbenzyloxy, and 3-trifluoromethylthiophenoxy;

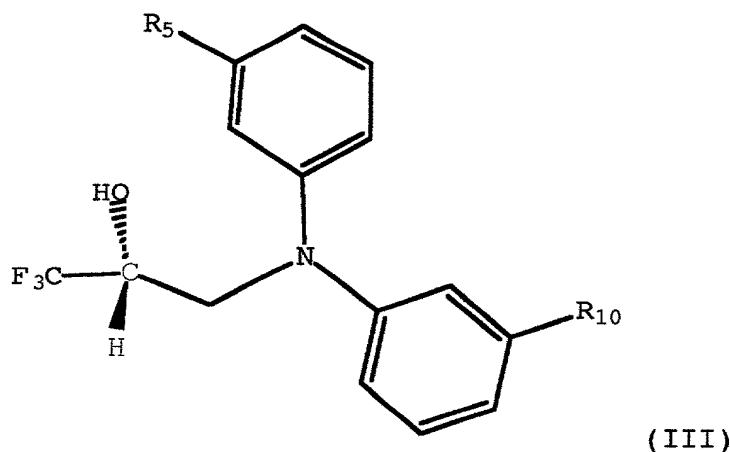
R_{10} is selected from the group consisting of 1,1,2,2-tetrafluoroethoxy,

pentafluoroethyl, and trifluoromethyl;

R_6 , R_7 , R_{11} , and R_{12} are independently hydrido or fluoro.

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94. The method of Claim 86, wherein said compound is a compound of Formula III:



wherein R_5 and R_{10} are selected to form a compound selected from the group

20 consisting of;

R_5 is 3-isopropylphenoxy and R_{10} is pentafluoroethyl;

R_5 is 2,3-dichlorophenoxy and R_{10} is pentafluoroethyl;

- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluorophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-methylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is pentafluoroethyl;
- 5 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-ethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-dimethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-*t*-butylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is pentafluoroethyl;
- 10 R_5 is 3,4-dichlorophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-dimethylaminophenoxy and R_{10} is pentafluoroethyl;
- 15 R_5 is 3-cyclopropylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-aminophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is pentafluoroethyl;
- 20 R_5 is 4-propoxyphenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is pentafluoroethyl;

- R_5 is 2-nitrophenoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is pentafluoroethyl;
- 5 R_5 is cyclohexylmethylenoxy and R_{10} is pentafluoroethyl;
- R_5 is benzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-ethylbenzyloxy and R_{10} is pentafluoroethyl;
- 10 R_5 is isopropoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is pentafluoroethyl;
- R_5 is isopropylthio and R_{10} is pentafluoroethyl;
- R_5 is cyclopentoxy and R_{10} is pentafluoroethyl;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is pentafluoroethyl;
- 15 R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is pentafluoroethyl;
- R_5 is 4-isopropylbenzyloxy and R_{10} is pentafluoroethyl;
- 20 R_5 is 1-phenylethoxy and R_{10} is pentafluoroethyl;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is pentafluoroethyl;

- R_5 is 3-trifluoromethylphenyl and R_{10} is pentafluoroethyl;
- R_5 is 4-methoxyphenylamino and R_{10} is pentafluoroethyl;
- R_5 is 4-nitrophenylthio and R_{10} is pentafluoroethyl;
- R_5 is 3-isopropylphenoxy and R_{10} is trifluoromethyl;
- 5 R_5 is 2,3-dichlorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-fluorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-methylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is trifluoromethyl;
- 10 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-ethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-dimethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-*t*-butylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is trifluoromethyl;
- 15 R_5 is 3,4-dichlorophenoxy and R_{10} is trifluoromethyl;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is trifluoromethyl;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-dimethylaminophenoxy and R_{10} is trifluoromethyl;
- 20 R_5 is 3-cyclopropylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is trifluoromethyl;

- R_5 is 3-pentafluoroethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-aminophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 4-propoxyphenoxy and R_{10} is trifluoromethyl;
- 5 R_5 is 3-trifluoromethylphenoxy and R_{10} is trifluoromethyl;
- R_5 is 2-nitrophenoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is trifluoromethyl;
- 10 R_5 is cyclohexylmethylenoxy and R_{10} is trifluoromethyl;
- R_5 is benzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 4-ethylbenzyloxy and R_{10} is trifluoromethyl;
- 15 R_5 is isopropoxy and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is trifluoromethyl;
- R_5 is isopropylthio and R_{10} is trifluoromethyl;
- R_5 is cyclopentoxy and R_{10} is trifluoromethyl;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is trifluoromethyl;
- 20 R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3,4-dimethylbenzyloxy and R_{10} is trifluoromethyl;

- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 4-isopropylbenzyloxy and R_{10} is trifluoromethyl;
- R_5 is 1-phenylethoxy and R_{10} is trifluoromethyl;
- 5 R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is trifluoromethyl;
- R_5 is 3-trifluoromethylphenyl and R_{10} is trifluoromethyl;
- R_5 is 4-methoxyphenylamino and R_{10} is trifluoromethyl;
- R_5 is 4-nitrophenylthio and R_{10} is trifluoromethyl;
- R_5 is 3-isopropylphenoxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 2,3-dichlorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-fluorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-methylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is trifluoromethoxy;
- 15 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-ethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-dimethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-*t*-butylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is trifluoromethoxy;
- 20 R_5 is 3,4-dichlorophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is trifluoromethoxy;

- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-dimethylaminophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-cyclopropylphenoxy and R_{10} is trifluoromethoxy;
- 5 R_5 is 3-(2-furyl)phenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-aminophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 4-propoxyphenoxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 3-trifluoromethylphenoxy and R_{10} is trifluoromethoxy;
- R_5 is 2-nitrophenoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-difluorobenzyloxy and R_{10} is trifluoromethoxy;
- 15 R_5 is cyclohexylmethylenoxy and R_{10} is trifluoromethoxy;
- R_5 is benzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 4-ethylbenzyloxy and R_{10} is trifluoromethoxy;
- 20 R_5 is isopropoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylbenzyl and R_{10} is trifluoromethoxy;

- R_5 is isopropylthio and R_{10} is trifluoromethoxy;
- R_5 is cyclopentoxy and R_{10} is trifluoromethoxy;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is trifluoromethoxy;
- 5 R_5 is 3,4-dimethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 4-isopropylbenzyloxy and R_{10} is trifluoromethoxy;
- R_5 is 1-phenylethoxy and R_{10} is trifluoromethoxy;
- 10 R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is trifluoromethoxy;
- R_5 is 3-trifluoromethylphenyl and R_{10} is trifluoromethoxy;
- R_5 is 4-methoxyphenylamino and R_{10} is trifluoromethoxy;
- R_5 is 4-nitrophenylthio and R_{10} is trifluoromethoxy;
- R_5 is 3-isopropylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 2,3-dichlorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-methylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-fluoro-5-bromophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 20 R_5 is 4-chloro-3-ethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-ethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;

- R_5 is 3,5-dimethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-*t*-butylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluoro-3-methylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,4-dichlorophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 5 R_5 is 5,6,7,8-tetrahydro-2-naphthoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-(1,1,2,2-tetrafluoroethoxy)phenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-difluoromethoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-dimethylaminophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 10 R_5 is 3-cyclopropylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-(2-furyl)phenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-pentafluoroethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-aminophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3,4,5-trimethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 4-propoxyphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylphenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-nitrophenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethoxybenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 20 R_5 is 3,5-difluorobenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is cyclohexylmethylenoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is benzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;

- R_5 is 3,5-ditrifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-trifluoromethoxybenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-ethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is isopropoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 5 R_5 is 3-trifluoromethylbenzyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is isopropylthio and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is cyclopentoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-chloro-5-pyridinyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylthiobenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 10 R_5 is 3,4-dimethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 2-fluoro-3-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-fluoro-5-trifluoromethylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- 15 R_5 is 4-isopropylbenzyloxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 1-phenylethoxy and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-fluoro-3-methylbenzoyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 3-trifluoromethylphenyl and R_{10} is 1,1,2,2-tetrafluoroethoxy;
- R_5 is 4-methoxyphenylamino and R_{10} is 1,1,2,2-tetrafluoroethoxy; and
- 20 R_5 is 4-nitrophenylthio and R_{10} is 1,1,2,2-tetrafluoroethoxy.

95. The method of Claim 86 further characterized by treating coronary artery disease in a subject by administering a therapeutically effective

amount of a compound of Claim 86 or a pharmaceutically acceptable salt thereof.

5 96. The method of Claim 86 further characterized by preventing coronary artery disease in a subject by administering a therapeutically effective amount of a compound of Claim 86 or a pharmaceutically acceptable salt thereof.

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 97. The method of Claim 86 further characterized by preventing cerebral vascular accident (CVA) in a subject by administering a therapeutically effective amount of a compound of Claim 86 or a pharmaceutically acceptable salt thereof.

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 98. The method of Claim 86 further characterized by treating or preventing dyslipidemia in a subject by administering a therapeutically effective amount of a compound of Claim 86 or a pharmaceutically acceptable salt thereof.

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**(R)-Chiral Halogenated Substituted N,N-Bis-Phenyl
Aminoalcohol Compounds Useful for Inhibiting Cholesteryl Ester
Transfer Protein Activity**

5

Abstract

The invention relates to substituted aryl and heteroaryl (R)-Chiral Halogenated 1-Substitutedamino-(n+1)-Alkanol compounds useful as inhibitors of cholesteryl ester transfer protein (CETP; plasma lipid transfer protein-I) and compounds, compositions and methods for treating atherosclerosis and other coronary artery diseases. Novel high yield, stereoselective processes for the preparation of the chiral substituted alkanol compounds from chiral and achiral intermediates are described. Preferred (R)-Chiral 1-Substitutedamino-(n+1)-Alkanol compounds are substituted (R)-Chiral N,N-bis-phenyl aminoalcohols. A preferred specific (R)-Chiral N,N-bis-phenyl aminoalcohol is the compound:

